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#46

RAW SEQUENCE LISTING

PATENT APPLICATION: US/08/803,954B

DATE: 02/27/2003 **%-6**TIME: 10:29:50

Input Set : A:\68430009.app

Output Set: N:\CRF4\02272003\H803954B.raw

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5 <110> APPLICANT: Langley, Keith E
        Boone, Thomas C
         DeClerck, Yves A
 9
13 <120> TITLE OF INVENTION: METALLOPROTEINASE INHIBITOR
17 <130> FILE REFERENCE: 06843.0009-08000
21 <140> CURRENT APPLICATION NUMBER: 08/803,954B
23 <141> CURRENT FILING DATE: 2001-08-29
27 <150> PRIOR APPLICATION NUMBER: 08/212,660
29 <151> PRIOR FILING DATE: 1994-03-11
33 <150> PRIOR APPLICATION NUMBER: 08/087,021
35 <151> PRIOR FILING DATE: 1993-07-06
39 <150> PRIOR APPLICATION NUMBER: 07/710,728
41 <151> PRIOR FILING DATE: 1991-06-03
45 <150> PRIOR APPLICATION NUMBER: 07/501,904
47 <151> PRIOR FILING DATE: 1990-03-29
51 <150> PRIOR APPLICATION NUMBER: 07/355,027
53 <151> PRIOR FILING DATE: 1989-05-19
57 <160> NUMBER OF SEQ ID NOS: 36
61 <170> SOFTWARE: PatentIn Ver. 2.0
65 <210> SEO ID NO: 1
67 <211> LENGTH: 13
69 <212> TYPE: PRT
71 <213> ORGANISM: Homo sapiens
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79 1
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85 <210> SEQ ID NO: 2
87 <211> LENGTH: 19
89 <212> TYPE: PRT
91 <213> ORGANISM: Homo sapiens
95 <400> SEQUENCE: 2
97 Cys Ser Cys Ser Pro Val His Pro Gln Gln Ala Phe Cys Asn Ala Asp
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103 Val Val Ile
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113 <211> LENGTH: 20
115 <212> TYPE: PRT
117 <213> ORGANISM: Homo sapiens
121 <400> SEQUENCE: 3
123 Val Val Phe Phe Pro Val Ala His Pro His Ser Trp Pro Thr Gln Val
125 1
                      5
                                         10
129 Ser Leu Arg Thr
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20

131

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Input Set : A:\68430009.app

Output Set: N:\CRF4\02272003\H803954B.raw

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137 <210> SEQ ID NO: 4
139 <211> LENGTH: 21
141 <212> TYPE: PRT
143 <213> ORGANISM: Homo sapiens
147 <400> SEQUENCE: 4
149 Val Val Gly Gly Pro Ala Val Ala His Pro His Ser Trp Pro Thr Gln
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151
     1
155 Val Ser Leu Arg Thr
157
163 <210> SEQ ID NO: 5
165 <211> LENGTH: 6
167 <212> TYPE: PRT
169 <213> ORGANISM: Homo sapiens
173 <400> SEQUENCE: 5
175 Cys Ser Cys Ser Pro Val
177
183 <210> SEQ ID NO: 6
185 <211> LENGTH: 1045
187 <212> TYPE: DNA
189 <213> ORGANISM: Bos taurus
193 <400> SEQUENCE: 6
195 aatteegget tetatggage actegggace aggteegegg egegegeact egetegeteg 60
197 cegececca gecagetete getteegege egecageege geceegegee teetegetge 120
199 accccgcgac ctagagccaa gaaagtttgt gtggcgagtg agggccggag aggagagcgc 180
201 gcccgcggag tgccgtccag accagcgcgg ccccggcgga gaggggagcg ccccgagccc 240
203 aggeggegge ggetageeeg agteegegae eeeegeeet eegeeegea tgggegeege 300
205 egecegeage etgeegeteg egttetgeet eetgetgetg gggaegetge teeeceggge 360
207 cgacgcctgc agctgctccc cggtgcaccc gcaacaggcg ttttgcaatg cagacatagt 420
209 gatcagggcc aaagcagtca ataagaagga ggtggactct ggcaacgaca tctacggcaa 480
211 ccccatcaag cggattcagt atgagatcaa gcagataaag atgttcaagg gacctgatca 540
213 ggacatagag tttatctaca cageceeege egetgeegtg tgtggggtet egetggacat 600
215 tggaggaaag aaggagtatc tcattgcagg gaaggccgag gggaatggca atatgcatat 660
217 caccetetgt gactteateg tgeeetggga caccetgagt gecacceaga agaagageet 720
219 gaaccacagg taccagatgg gctgtgagtg caagatcact cgatgcccca tgatcccatg 780
221 ctacatetee teteeggaeg agtgeetetg gatggaetgg gteaeggaga agaacateaa 840
223 eggacaceag gecaagttet tegeetgeat caagagaage gaeggeteet gegeetggta 900
225 ccgcggagca gcaccccca agcaggagtt tctggacatg aggacccgta agcaggccac 960
227 caggactect ggggecaatt gacagtgtee aagagtteag actggteeag eteegacate 1020
229 ccttcctgga cacagcatga ataaa
233 <210> SEQ ID NO: 7
235 <211> LENGTH: 220
237 <212> TYPE: PRT
239 <213> ORGANISM: Bos taurus
243 <400> SEQUENCE: 7
245 Met Gly Ala Ala Ala Arg Ser Leu Pro Leu Ala Phe Cys Leu Leu Leu
247
                                         10
251 Leu Gly Thr Leu Leu Pro Arg Ala Asp Ala Cys Ser Cys Ser Pro Val
                 20
                                     25
257 His Pro Gln Gln Ala Phe Cys Asn Ala Asp Ile Val Ile Arg Ala Lys
```

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PATENT APPLICATION: US/08/803,954B TIME: 10:29:50

Input Set : A:\68430009.app

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```
259
             35
                                 40
263 Ala Val Asn Lys Lys Glu Val Asp Ser Gly Asn Asp Ile Tyr Gly Asn
                             55
269 Pro Ile Lys Arg Ile Gln Tyr Glu Ile Lys Gln Ile Lys Met Phe Lys
271 65
                         70
275 Gly Pro Asp Gln Asp Ile Glu Phe Ile Tyr Thr Ala Pro Ala Ala Ala
                     85
                                         90
281 Val Cys Gly Val Ser Leu Asp Ile Gly Gly Lys Lys Glu Tyr Leu Ile
                                    105
287 Ala Gly Lys Ala Glu Gly Asn Gly Asn Met His Ile Thr Leu Cys Asp
289
                                120
            115
293 Phe Ile Val Pro Trp Asp Thr Leu Ser Ala Thr Gln Lys Lys Ser Leu
295
                            135
                                                140
299 Asn His Arg Tyr Gln Met Gly Cys Glu Cys Lys Ile Thr Arg Cys Pro
                                            155
                        150
305 Met Ile Pro Cys Tyr Ile Ser Ser Pro Asp Glu Cys Leu Trp Met Asp
307
                                        170
                    165
311 Trp Val Thr Glu Lys Asn Ile Asn Gly His Gln Ala Lys Phe Phe Ala
313
               180
                                    185
317 Cys Ile Lys Arg Ser Asp Gly Ser Cys Ala Trp Tyr Arg Gly Ala Ala
                                200
           195
323 Pro Pro Lys Gln Glu Phe Leu Asp Ile Glu Asp Pro
        210
331 <210> SEQ ID NO: 8
333 <211> LENGTH: 1032
335 <212> TYPE: DNA
337 <213> ORGANISM: Homo sapiens
341 <400> SEQUENCE: 8
343 atteeggeee geegteeece acceegeege eeegeeegge gaattgegee eegegeeeet 60
345 cccctcgcgc ccccgagaca aagaggagag aagtttgcgc ggccgagcgg ggcaggtgag 120
347 gagggtgage egegegggag gggeeegeet eggeeeegge teageeeeeg eeegegeeee 180
349 cagecegeeg cegegageag egeeeggaee eeceagegge ggeeeeegee egeeeageee 240
351 cooggeooge catgggegee geggeoogea cootgegget ggegetegge etectgetge 300
353 tggcgacgct gcttcgcccg gccgacgcct gcagctgctc cccggtgcac ccgcaacagg 360
355 cgttttgcaa tgcagatgta gtgatcaggg ccaaagcggt cagtgagaag gaagtggact 420
357 ctggaaacga catttatggc aaccctatca agaggatcca gtatgagatc aagcagataa 480
359 agatgttcaa agggcctgag aaggatatag agtttatcta cacggccccc tcctcggcag 540
361 tqtqtqqqqt ctcqctqqac gttgqaqgaa agaaqgaata tctcattgca ggaaaggccg 600
363 agggggacgg caagatgcac atcaccctct gtgacttcat cgtgccctgg gacaccctga 660
365 gcaccaccca gaagaagage etgaaccaca ggtaccagat gggetgegag tgcaagatea 720
367 egegetgeee catgateeeg tgetacatet eeteeeegga egagtgeete tggatggaet 780
369 gggtcacaga gaagaacatc aacgggcacc aggccaagtt cttcgcctgc atcaagagaa 840
371 gtgacggctc ctgtgcgtgg taccgcggcg cggcgccccc caagcaggag tttctcgaca 900
373 tegaggaece ataageagge etecaaegee eetgtggeea aetgeaaaaa aageeteeaa 960
375 gggtttegae tggteeaget etgaeateee tteetggaaa eageatgaat aaaacaetea 1020
377 tccccggaat tc
                                                                       1032
381 <210> SEQ ID NO: 9
383 <211> LENGTH: 220
385 <212> TYPE: PRT
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RAW SEQUENCE LISTING DATE: 02/27/2003 PATENT APPLICATION: US/08/803,954B TIME: 10:29:50

Input Set : A:\68430009.app

Output Set: N:\CRF4\02272003\H803954B.raw

```
387 <213> ORGANISM: Homo sapiens
391 <400> SEQUENCE: 9
393 Met Gly Ala Ala Ala Arg Thr Leu Arg Leu Ala Leu Gly Leu Leu
                                         10
399 Leu Ala Thr Leu Leu Arg Pro Ala Asp Ala Cys Ser Cys Ser Pro Val
               20
                                    25
405 His Pro Gln Gln Ala Phe Cys Asn Ala Asp Val Val Ile Arg Ala Lys
411 Ala Val Ser Glu Lys Glu Val Asp Ser Gly Asn Asp Ile Tyr Gly Asn
        50
417 Pro Ile Lys Arg Ile Gln Tyr Glu Ile Lys Gln Ile Lys Met Phe Lys
419 65
                        70
423 Gly Pro Glu Lys Asp Ile Glu Phe Ile Tyr Thr Ala Pro Ser Ser Ala
429 Val Cys Gly Val Ser Leu Asp Val Gly Gly Lys Lys Glu Tyr Leu Ile
                                   105
               100
435 Ala Gly Lys Ala Glu Gly Asp Gly Lys Met His Ile Thr Leu Cys Asp
           115
                               120
441 Phe Ile Val Pro Trp Asp Thr Leu Ser Thr Thr Gln Lys Lys Ser Leu
                           135
                                               140
443 130
447 Asn His Arg Tyr Gln Met Gly Cys Glu Cys Lys Ile Thr Arg Cys Pro
                                           155
                       150
453 Met Ile Pro Cys Tyr Ile Ser Ser Pro Asp Glu Cys Leu Trp Met Asp
                   165
                                        170
459 Trp Val Thr Glu Lys Asn Ile Asn Gly His Gln Ala Lys Phe Phe Ala
                                   185
465 Cys Ile Lys Arg Ser Asp Gly Ser Cys Ala Trp Tyr Arg Gly Ala Ala
                               200
467 195
471 Pro Pro Lys Gln Glu Phe Leu Asp Ile Glu Asp Pro
473 210
                           215
479 <210> SEQ ID NO: 10
481 <211> LENGTH: 45
483 <212> TYPE: PRT
485 <213> ORGANISM: Bos taurus
489 <400> SEQUENCE: 10
491 Cys Ser Cys Ser Pro Cys His Pro Gln Gln Ala Phe Cys Asn Ala Asp
                                        10
497 Ile Val Ile Arg Ala Lys Ala Val Asn Lys Lys Glu Val Asp Ser Gly
               20
                                    25
503 Asn Asp Ile Tyr Gly Asn Pro Ile Lys Arg Lys Gln Tyr
                                40
511 <210> SEQ ID NO: 11
513 <211> LENGTH: 49
515 <212> TYPE: PRT
517 <213> ORGANISM: Bos taurus
521 <400> SEQUENCE: 11
523 Cys Thr Cys Val Pro Pro His Pro Gln Thr Ala Phe Cys Asn Ser Asp
                    .5
                                        10
529 Val Val Ile Arg Ala Lys Phe Val Gly Thr Ala Glu Val Asn Glu Thr
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Output Set: N:\CRF4\02272003\H803954B.raw

```
25
                 20
531
535 Ala Leu Leu Tyr Arg Tyr Leu Ile Lys Met Leu Lys Met Pro Ser Gly
                                 40
541 Phe
549 <210> SEQ ID NO: 12
551 <211> LENGTH: 49
553 <212> TYPE: PRT
555 <213> ORGANISM: Homo sapiens
559 <400> SEQUENCE: 12
561 Cys Thr Cys Val Pro Pro His Pro Gln Thr Ala Phe Cys Asn Ser Asp
                                          10
567 Leu Val Ile Arg Ala Lys Phe Val Gly Thr Pro Glu Val Asn Gln Thr
573 Thr Leu Tyr Gln Arg Tyr Glu Ile Lys Met Thr Lys Met Tyr Lys Gly
575
                                 40
579 Phe
587 <210> SEQ ID NO: 13
589 <211> LENGTH: 49
591 <212> TYPE: PRT
593 <213> ORGANISM: Bos taurus
597 <400> SEQUENCE: 13
599 Cys Thr Cys Val Pro Pro His Pro Gln Thr Ala Phe Cys Asn Ser Asp
                                         10
605 Val Val Ile Arg Ala Lys Phe Val Gly Thr Ala Glu Val Asn Glu Thr
                 20
611 Ala Leu Leu Tyr Arg Tyr Leu Ile Lys Met Leu Lys Met Pro Ser Gly
613
                                 40
617 Phe
625 <210> SEQ ID NO: 14
627 <211> LENGTH: 45
629 <212> TYPE: PRT
631 <213> ORGANISM: Bos taurus
635 <400> SEQUENCE: 14
637 Cys Ser Cys Ser Pro Val His Pro Gln Gln Ala Phe Cys Asn Ala Asp
                                          10
643 Ile Val Ile Arg Ala Lys Ala Val Asn Lys Lys Glu Val Asp Ser Gly
                20
                                      25
649 Asn Asp Ile Tyr Gly Asn Pro Ile Lys Arg Ile Gln Tyr
             35
                                 40
657 <210> SEQ ID NO: 15
659 <211> LENGTH: 45
661 <212> TYPE: DNA
663 <213> ORGANISM: Bos taurus
667 <400> SEQUENCE: 15
669 gatcacaatg tcagcattgc agaaggcctg ctggggatgc acagg
                                                                       45
673 <210> SEQ ID NO: 16
675 <211> LENGTH: 30
677 <212> TYPE: DNA
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679 <213> ORGANISM: Artificial Sequence

RAW SEQUENCE LISTING ERROR SUMMARY DATE: 02/27/2003 PATENT APPLICATION: US/08/803,954B TIME: 10:29:51

Input Set : A:\68430009.app

Output Set: N:\CRF4\02272003\H803954B.raw

Please Note:

Use of n and/or Xaa have been detected in the Sequence Listing. Please review the Sequence Listing to ensure that a corresponding explanation is presented in the <220> to <223> fields of each sequence which presents at least one n or Xaa.

Seq#:16; N Pos. 4,19,22,28 Seq#:17; N Pos. 4,7,13,19,28

Invalid Line Length:

The rules require that a line not exceed 72 characters in length. This includes spaces.

```
Seq#:1; Line(s) 1,3,5,7,9,11,13,15,19,21,23,25,27,29,31,33,35,37,39,41,43
Seq#:1; Line(s) 45,47,49,51,53,55,57,59,61,63,65,67,69,71,73,75,77,79,81,83
Seq#:1; Line(s) 85
Seq#:2; Line(s) 87,89,91,93,95,97,99,101,103,105,107,109,111
Seq#:3; Line(s) 113,115,117,119,121,123,125,127,129,131,133,135,137
Seq#:4; Line(s) 139,141,143,145,147,149,151,153,155,157,159,161,163
Seq#:5; Line(s) 165,167,169,171,173,175,177,179,181,183
Seq#:6; Line(s) 185,187,189,191,193,195,197,199,201,203,205,207,209,211,213
Seg#:6; Line(s) 215,217,219,221,223,225,227,229,231,233
Seq#:7; Line(s) 235,237,239,241,243,245,247,249,251,253,255,257,259,261,263
Seq#:7; Line(s) 265,267,269,271,273,275,277,279,281,283,285,287,289,291,293
Seq#:7; Line(s) 295,297,299,301,303,305,307,309,311,313,315,317,319,321,323
Seq#:7; Line(s) 325,327,329,331
Seq#:8; Line(s) 333,335,337,339,341,343,345,347,349,351,353,355,357,359,361
Seq#:8; Line(s) 363,365,367,369,371,373,375,377,379,381
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Seq#:9; Line(s) 413,415,417,419,421,423,425,427,429,431,433,435,437,439,441
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Seq#:9; Line(s) 473,475,477,479
Seq#:10; Line(s) 481,483,485,487,489,491,493,495,497,499,501,503,505,507
Seq#:10; Line(s) 509,511
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Seq#:13; Line(s) 617,619,621,623,625
Seq#:14; Line(s) 627,629,631,633,635,637,639,641,643,645,647,649,651,653
Seq#:14; Line(s) 655,657
Seq#:15; Line(s) 659,661,663,665,667,669,671,673
Seq#:16; Line(s) 675,677,679,681,683,685,687,689,691,693,695,697,699,701
Seq#:16; Line(s) 703,705,707,709,711,713,715,717,719,721,723,725,727,729
Seq#:16; Line(s) 731,733,735,737
Seq#:17; Line(s) 739,741,743,745,747,749,751,753,755,757,759,761,763,765
Seq#:17; Line(s) 767,769,771,773,775,777,779,781,783,785,787,789,791,793
Seq#:17; Line(s) 795,797,799,801,803,805,807,809,811
Seq#:18; Line(s) 813,815,817,819,821,823,825,827
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RAW SEQUENCE LISTING ERROR SUMMARY DATE: 02/27/2003 PATENT APPLICATION: US/08/803,954B TIME: 10:29:51

Input Set : A:\68430009.app

Output Set: N:\CRF4\02272003\H803954B.raw

Seq#:19; Line(s) 829,831,833,835,837,839,841,843
Seq#:20; Line(s) 845,847,849,851,853,855,857,859
Seq#:21; Line(s) 861,863,865,867,869,871,873,875
Seq#:22; Line(s) 877,879,881,883,885,887,889,891,893,895,897,899
Seq#:23; Line(s) 901,903,905,907,909,911,913,915,917,919,921,923
Seq#:24; Line(s) 925,927,929,931,933,935,937,939,941,943,945,947,949,951
Seq#:25; Line(s) 953,955,957,959,961,963,965,967,969,971,973,975,977,979
Seq#:26; Line(s) 981,983,985,987,989,991,993,995,997,999,1001,1003,1005
Seq#:27; Line(s) 1007
Seq#:27; Line(s) 1009,1011,1013,1015,1017,1019,1021,1023,1025,1027,1029
Seq#:27; Line(s) 1031
Seq#:28; Line(s) 1033,1035,1037,1039,1041,1043,1045,1047,1049,1051,1053
Seq#:28; Line(s) 1055
Seq#:29; Line(s) 1057,1059,1061,1063,1065,1067,1069,1071,1073,1075,1077

VERIFICATION SUMMARY

DATE: 02/27/2003

PATENT APPLICATION: US/08/803,954B

TIME: 10:29:51

Input Set : A:\68430009.app
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L:733 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:16 after pos.:0 L:807 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:17 after pos.:0